

PTO-1390 (Rev. 07-2005)

Approved for use through 03/31/2007. OMB 0651-0021  
U. S. Patent and Trademark Office; U.S. DEPARTMENT OF COMMERCE

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<b>TRANSMITTAL LETTER TO THE UNITED STATES DESIGNATED/ELECTED OFFICE (DO/EO/US) CONCERNING A SUBMISSION UNDER 35 U.S.C. 371</b>		ATTORNEY'S DOCKET NUMBER 65645(46590)
		U.S. APPLICATION NO. (if known, see 37 CFR 1.5) <b>10/581008</b>
INTERNATIONAL APPLICATION NO. PCT/JP2004/017995	INTERNATIONAL FILING DATE 26 November 2004	PRIORITY DATE CLAIMED 27 November 2003
TITLE OF INVENTION METHOD OF ESTIMATING TOXICITY OF DRUG		
APPLICANT(S) FOR DO/EO/US Kenji Takami et al.		
Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information:		
<p>1. <input checked="" type="checkbox"/> This is a <b>FIRST</b> submission of items concerning a submission under 35 U.S.C. 371.</p> <p>2. <input type="checkbox"/> This is a <b>SECOND</b> or <b>SUBSEQUENT</b> submission of items concerning a submission under 35 U.S.C. 371.</p> <p>3. <input type="checkbox"/> This is an express request to begin national examination procedures (35 U.S.C. 371(f)). The submission must include items (5), (6), (9) and (21) indicated below.</p> <p>4. <input type="checkbox"/> The US has been elected (Article 31).</p> <p>5. <input checked="" type="checkbox"/> A copy of the International Application as filed (35 U.S.C. 371 (c)(2))</p> <p style="margin-left: 20px;">a. <input type="checkbox"/> is attached hereto (required only if not communicated by the International Bureau).</p> <p style="margin-left: 20px;">b. <input checked="" type="checkbox"/> has been communicated by the International Bureau.</p> <p style="margin-left: 20px;">c. <input type="checkbox"/> is not required, as the application was filed in the United States Receiving Office (RO/US).</p> <p>6. <input checked="" type="checkbox"/> An English language translation of the International Application as filed (35 U.S.C. 371(c)(2)).</p> <p style="margin-left: 20px;">a. <input checked="" type="checkbox"/> is attached hereto.</p> <p style="margin-left: 20px;">b. <input type="checkbox"/> has been previously submitted under 35 U.S.C. 154(d)(4).</p> <p>7. <input type="checkbox"/> Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371(c)(3))</p> <p style="margin-left: 20px;">a. <input type="checkbox"/> are attached hereto (required only if not communicated by the International Bureau).</p> <p style="margin-left: 20px;">b. <input type="checkbox"/> have been communicated by the International Bureau.</p> <p style="margin-left: 20px;">c. <input type="checkbox"/> have not been made; however, the time limit for making such amendments has NOT expired.</p> <p style="margin-left: 20px;">d. <input type="checkbox"/> have not been made and will not be made.</p> <p>8. <input type="checkbox"/> An English language translation of the amendments to the claims under PCT Article 19 (35 U.S.C. 371(c)(3)).</p> <p>9. <input type="checkbox"/> An oath or declaration of the inventor(s) (35 U.S.C. 371(c)(4)).</p> <p>10. <input type="checkbox"/> An English language translation of the annexes of the International Preliminary Examination Report under PCT Article 36 (35 U.S.C. 371(c)(5)).</p> <p><b>Items 11 to 20 below concern document(s) or information included:</b></p> <p>11. <input type="checkbox"/> An Information Disclosure Statement under 37 CFR 1.97 and 1.98.</p> <p>12. <input type="checkbox"/> An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included.</p> <p>13. <input type="checkbox"/> A preliminary amendment.</p> <p>14. <input checked="" type="checkbox"/> An Application Data Sheet under 37 CFR 1.76.</p> <p>15. <input type="checkbox"/> A substitute specification.</p> <p>16. <input type="checkbox"/> A power of attorney and/or change of address letter.</p> <p>17. <input checked="" type="checkbox"/> A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 37 CFR 1.821 – 1.825.</p> <p>18. <input type="checkbox"/> A second copy of the published International Application under 35 U.S.C. 154(d)(4).</p> <p>19. <input type="checkbox"/> A second copy of the English language translation of the international application under 35 U.S.C. 154(d)(4).</p>		

1AP20 Rec'd PCT/PTO 26 MAY 2006

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U.S. APPLICATION NO. <b>10/591008</b> (If known, see 37 CFR 1.51)	INTERNATIONAL APPLICATION NO. <b>PCT/JP2004/017995</b>	ATTORNEY'S DOCKET NUMBER <b>65645(46590)</b>	
20. <input checked="" type="checkbox"/> Other items or information: Figures 1 - 7 (7 sheets); Sequence Listing (44 pages); Sequence Diskette; Sequence Statement (2 pages); English Translation of PCT Request (5 pages); Copy of Cover Page of Published PCT Application No. WO 2005/052154 A1; Copy of International Search Report (Form PCT/ISA/210) (3 pages); Copy of Form PCT/IB/301, PCT/IB/304, & PCT/IB/308; Return Receipt Postcard.			
The following fees have been submitted		<b>CALCULATIONS</b>	<b>PTO USE ONLY</b>
21. <input checked="" type="checkbox"/> Basic national fee (37 CFR 1.492(a)) ..... \$300		\$ 300.00	
22. <input checked="" type="checkbox"/> Examination fee (37 CFR 1.492(c)) If the written opinion prepared by ISA/US or the international preliminary examination report prepared by IPEA/US indicates all claims satisfy provisions of PCT Article 33(1)-(4) ..... \$0 All other situations ..... \$200		\$ 200.00	
23. <input checked="" type="checkbox"/> Search fee (37 CFR 1.492(b)) If the written opinion of the ISA/US or the international preliminary examination report prepared by IPEA/US indicates all claims satisfy provisions of PCT Article 33(1)-(4) ..... \$0 Search fee (37 CFR 1.445(a)(2)) has been paid on the international application to the USPTO as an International Searching Authority ..... \$100 International Search Report prepared by an ISA other than the US and provided to the Office or previously communicated to the US by the IB ..... \$400 All other situations ..... \$500		\$ 400.00	
<b>TOTAL OF 21, 22 and 23 =</b>		<b>\$ 900.00</b>	
<input type="checkbox"/> Additional fee for specification and drawings filed in paper over 100 sheets (excluding sequence listing in compliance with 37 CFR 1.821(c) or (e) or computer program listing in an electronic medium) (37 CFR 1.492(j)). The fee is \$250 for each additional 50 sheets of paper or fraction thereof.			
Total Sheets	Extra Sheets	Number of each additional 50 or fraction thereof (round up to a whole number)	RATE
61 - 100 =	/50 =		x \$250.00
			\$
Surcharge of \$130 for furnishing any of the search fee, examination fee, or the oath or declaration after the date of commencement of the national stage (37 CFR 1.492(h)).		\$	
CLAIMS	NUMBER FILED	NUMBER EXTRA	RATE
Total claims	13 - 20 =	x	0.00
Independent claims	5 - 3 =	2 x	200.00
MULTIPLE DEPENDENT CLAIM(S) (if applicable)		+	360.00
<b>TOTAL OF ABOVE CALCULATIONS =</b>		<b>\$ 1,660.00</b>	
<input type="checkbox"/> Applicant claims small entity status. See 37 CFR 1.27. Fees above are reduced by 1/2.			
<b>SUBTOTAL =</b>		<b>\$ 1,660.00</b>	
Processing fee of \$130.00 for furnishing the English translation later than 30 months from the earliest claimed priority date (37 CFR 1.492(i)).		\$	
<b>TOTAL NATIONAL FEE =</b>		<b>\$ 1,660.00</b>	
Fee for recording the enclosed assignment (37 CFR 1.21(h)). The assignment must be accompanied by an appropriate cover sheet (37 CFR 3.28, 3.31). \$40.00 per property		\$	
		\$	
<b>TOTAL FEES ENCLOSED =</b>		<b>\$ 1,660.00</b>	
		Amount to be refunded:	\$
		Amount to be charged	\$

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- a. ☐ A check in the amount of \$ \_\_\_\_\_ to cover the above fees is enclosed.
- b. ☒ Please charge my Deposit Account No. 04-1105 in the amount of \$ 1,660.00 to cover the above fees.  
A duplicate copy of this sheet is enclosed.
- c. ☒ The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any overpayment to Deposit Account No. 04-1105. A duplicate copy of this sheet is enclosed.
- d. ☐ Fees are to be charged to a credit card. **WARNING:** Information on this form may become public. Credit card information should not be included on this form. Provide credit card information and authorization on PTO-2038

NOTE: Where an appropriate time limit under 37 CFR 1.495 has not been met, a petition to revive (37 CFR 1.137(a) or (b)) must be filed and granted to restore the International Application to pending status.

SEND ALL CORRESPONDENCE TO:

David G. Conlin  
EDWARDS ANGELL PALMER & DODGE LLP  
P.O. Box 55874  
Boston, Massachusetts 02205  
(617) 439-4444

SIGNATURE

Jonathan M. Sparks, Ph.D.

NAME

CUSTOMER NUMBER: 21874

53,624

REGISTRATION NUMBER

AP20 Rec'd PCT/PTO 26 MAY 2006

PTO/SB/17 (01-06)

Approved for use through 7/31/2006. OMB 0651-0032

U.S. Patent and Trademark Office; U.S. DEPARTMENT OF COMMERCE

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<b>FEE TRANSMITTAL</b> <b>For FY 2006</b>		<b>Complete if Known</b> Application Number <b>Not Yet Assigned</b> Filing Date <b>Concurrently Herewith</b> First Named Inventor <b>Kenji Takami</b> Examiner Name <b>Not Yet Assigned</b> Art Unit <b>N/A</b> Attorney Docket No. <b>65645(46590)</b>	
<input type="checkbox"/> Applicant claims small entity status. See 37 CFR 1.27			
<b>TOTAL AMOUNT OF PAYMENT</b> (\$)    1,660.00			

**METHOD OF PAYMENT** (check all that apply)

☐ Check    ☐ Credit Card    ☐ Money Order    ☐ None    ☐ Other (please identify): \_\_\_\_\_

☒ Deposit Account    Deposit Account Number: **04-1105**    Deposit Account Name: **Edwards Angell Palmer & Dodge LLP**

For the above-identified deposit account, the Director is hereby authorized to: (check all that apply)

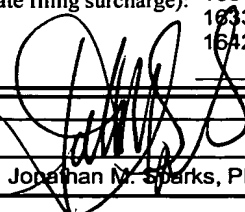
☒ Charge fee(s) indicated below                      ☐ Charge fee(s) indicated below, except for the filing fee  
☒ Charge any additional fee(s) or underpayment of fee(s) under 37.CFR 1.16 and 1.17                      ☒ Credit any overpayments

**FEE CALCULATION (All the fees below are due upon filing or may be subject to a surcharge.)**

Application Type	FILING FEES		SEARCH FEES		EXAMINATION FEES		Fees Paid (\$)
	Fee (\$)	Small Entity Fee (\$)	Fee (\$)	Small Entity Fee (\$)	Fee (\$)	Small Entity Fee (\$)	
Utility	300	150	500	250	200	100	
Design	200	100	100	50	130	65	
Plant	200	100	300	150	160	80	
Reissue	300	150	500	250	600	300	
Provisional	200	100	0	0	0	0	

2. EXCESS CLAIM FEES				Small Entity	
<u>Fee Description</u>				<u>Fee (\$)</u>	<u>Fee (\$)</u>
Each claim over 20 (including Reissues)				50	25
Each independent claim over 3 (including Reissues)				200	100
Multiple dependent claims				360	180
<u>Total Claims</u>	<u>Extra Claims</u>	<u>Fee (\$)</u>	<u>Fee Paid (\$)</u>	<u>Multiple Dependent Claims</u>	
<u>13</u>	- 20 =	x	=	<u>Fee (\$)</u>	<u>Fee Paid (\$)</u>
HP = highest number of total claims paid for, if greater than 20.				<u>360.00</u>	<u>360.00</u>
<u>Indep. Claims</u>	<u>Extra Claims</u>	<u>Fee (\$)</u>	<u>Fee Paid (\$)</u>		
<u>5</u>	- 3 =	x	=		
HP = highest number of independent claims paid for, if greater than 3.					

3. APPLICATION SIZE FEE		Fee (\$)	Fee Paid (\$)
If the specification and drawings exceed 100 sheets of paper (excluding electronically filed sequence or computer listings under 37 CFR 1.52(e)), the application size fee due is \$250 (\$125 for small entity) for each additional 50 sheets or fraction thereof. See 35 U.S.C. 41(a)(1)(G) and 37 CFR 1.16(s).			
<b>Total Sheets</b> <b>Extra Sheets</b> <b>Number of each additional 50 or fraction thereof</b> <b>Fee (\$)</b> <b>Fee Paid (\$)</b>			
61    - 100 =    /50    (round up to a whole number) x    =			
4. OTHER FEE(S)		Fees Paid (\$)	
Non-English Specification, \$130 fee (no small entity discount)			
Other (e.g., late filing surcharge):			
1631 Basic National Stage fee		300.00	
1633 National Stage Examination Fee - all other ...		200.00	
1642 National Stage Search Fee - search report ...		400.00	

<b>SUBMITTED BY</b>		Registration No.	Telephone
Signature		53,624	(617) 439-4444
Name (Print/Type)	Jonathan M. Sparks, Ph.D.	Date	May 26, 2006

00/581008

AP20 Rec'd PCT/PTO 26 MAY 2006

Application No. (if known): Not Yet Assigned

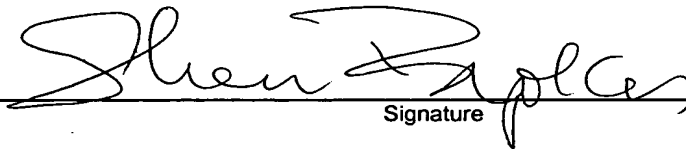
Attorney Docket No.: 65645(46590)

## Certificate of Express Mailing Under 37 CFR 1.10

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Sharon Bizokas

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Registration Number, if applicable

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Transmittal Letter to the United States Designated-Elected Office (3 pages)  
Fee Transmittal (1 page)  
Application Data Sheet (  
Specification excluding claims (50 pages), Claims (3 pages), Abstract (1 page)  
Figures 1 - 7 (7 sheets)  
Sequence Listing (44 pages)  
Sequence Diskette  
Sequence Statement (2 pages)  
English Translation of PCT Request (5 pages)  
Copy of Cover Page of Published PCT Application No. WO 2005/052154 A1 (1 page)  
Copy of International Search Report (Form PCT/ISA/210) (3 pages)  
Copy of Form PCT/IB/301, PCT/IB/304, & PCT/IB/308 (5 pages total)  
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